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## **CLAIMS**

## What is claimed is:

- 1. A method of treating a cognitive deficit associated with a central nervous system disorder or condition in an animal in need of said treatment comprising the steps of:
  - administering to said animal an augmenting agent which enhances CREB (a) pathway function; and
  - training said animal under conditions sufficient to produce an (b) improvement in performance by said animal of a cognitive task whose deficit is associated with said central nervous system disorder or condition.

whereby said cognitive deficit is treated.

- The method of Claim 1 wherein said animal has undergone neuronal stem cell 2. manipulation.
- 15 3. The method of Claim 1 wherein a performance gain is achieved relative to the performance of said cognitive task achieved by training alone.
  - 4. The method of Claim 1 wherein in step b), training comprises multiple training sessions.
- 5. The method of Claim 4 wherein said augmenting agent is administered before 20 and during each training session.
  - 6. The method of Claim 1 wherein said animal is a mammal.

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- 7. The method of Claim 6 wherein said mammal is a human.
- 8. The method of Claim 1 wherein said augmenting agent induces CREB-dependent gene expression.
- 9. The method of Claim 8 wherein said augmenting agent up-regulates a positive effector of CREB pathway function.
  - 10. The method of Claim 9 wherein said positive effector of CREB pathway function is a CREB activator.
  - 11. The method of Claim 8 wherein said augmenting agent down-regulates a negative effector of CREB pathway function.
- 10 12. The method of Claim 11 wherein said negative effector of CREB pathway function is a CREB repressor.
  - 13. The method of Claim 1 wherein said augmenting agent is a CREB functional modulator.
- 14. A method of enhancing a specific aspect of cognitive performance in an animal in need thereof comprising the steps of:
  - (a) administering to said animal an augmenting agent which enhances CREB pathway function; and
  - (b) training said animal under conditions sufficient to produce an improvement in performance of a specified cognitive task by said animal, whereby said specific aspect of cognitive performance is enhanced.

- 15. The method of Claim 14 wherein a performance gain is achieved relative to the performance of said specified cognitive task achieved by training alone.
- 16. The method of Claim 14 wherein in step b), training comprises multiple training sessions.
- 5 17. The method of Claim 16 wherein said augmenting agent is administered before and during each training session.
  - 18. The method of Claim 14 wherein said animal is a mammal.
  - 19. The method of Claim 18 wherein said mammal is a human.
- The method of Claim 14 wherein said augmenting agent induces CREB dependent gene expression.
  - 21. The method of Claim 20 wherein said augmenting agent up-regulates a positive effector of CREB pathway function.
  - 22. The method of Claim 21 wherein said positive effector of CREB pathway function is a CREB activator.
- 15 23. The method of Claim 20 wherein said augmenting agent down-regulates a negative effector of CREB pathway function.
  - 24. The method of Claim 23 wherein said negative effector of CREB pathway function is a CREB repressor.
- The method of Claim 14 wherein said augmenting agent is a CREB functionalmodulator.

- 26. A method of treating a cognitive deficit associated with age-associated memory impairment in an animal in need of said treatment comprising the steps of:
  - (a) administering to said animal an augmenting agent which enhances CREB pathway function; and
  - (b) training said animal under conditions sufficient to produce an improvement in performance by said animal of a cognitive task whose deficit is associated with age-associated memory impairment, whereby said cognitive deficit is treated.
- The method of Claim 26 wherein a performance gain is achieved relative to the performance of said cognitive task achieved by training alone.
  - 28. The method of Claim 26 wherein in step b), training comprises multiple training sessions.
  - 29. The method of Claim 28 wherein said augmenting agent is administered before and during each training session.
- 15 30. The method of Claim 26 wherein said animal is a mammal.
  - 31. The method of Claim 30 wherein said mammal is a human.
  - 32. The method of Claim 26 wherein said augmenting agent induces CREB-dependent gene expression.
- A method of treating a cognitive deficit associated with a neurodegenerative disease in an animal in need of said treatment comprising the steps of:
  - (a) administering to said animal an augmenting agent which enhances CREB pathway function; and

- (b) training said animal under conditions sufficient to produce an improvement in performance by said animal of a cognitive task whose deficit is associated with said neurodegenerative disease, whereby said cognitive deficit is treated.
- 5 34. The method of Claim 33 wherein a performance gain is achieved relative to the performance of said cognitive task achieved by training alone.
  - 35. The method of Claim 33 wherein said neurodegenerative disease is selected from the group consisting of: delirium, dementia, Alzheimer's disease, Parkinson's disease and Huntington's disease.
- 10 36. The method of Claim 33 wherein in step b), training comprises multiple training sessions.
  - 37. The method of Claim 36 wherein said augmenting agent is administered before and during each training session.
  - 38. The method of Claim 33 wherein said animal is a mammal.
- 15 39. The method of Claim 38 wherein said mammal is a human.
  - 40. The method of Claim 33 wherein said augmenting agent induces CREB-dependent gene expression.
  - 41. A method of treating a cognitive deficit associated with a psychiatric disease in an animal in need of said treatment comprising the steps of:
- 20 (a) administering to said animal an augmenting agent which enhances CREB pathway function; and

- (b) training said animal under conditions sufficient to produce an improvement in performance by said animal of a cognitive task whose deficit is associated with said psychiatric disease, whereby said cognitive deficit is treated.
- 5 42. The method of Claim 41 wherein a performance gain is achieved relative to the performance of said cognitive task achieved by training alone.
  - 43. The method of Claim 41 wherein said psychiatric disease is selected from the group consisting of: depression, schizophrenia, autism and attention deficit disorder.
- 10 44. The method of Claim 41 wherein in step b), training comprises multiple training sessions.
  - 45. The method of Claim 44 wherein said augmenting agent is administered before and during each training session.
  - 46. The method of Claim 41 wherein said animal is a mammal.
- 15 47. The method of Claim 46 wherein said mammal is a human.
  - 48. The method of Claim 41 wherein said augmenting agent induces CREB-dependent gene expression.
  - 49. A method of treating a cognitive deficit associated with cerebrovascular disease in an animal in need of said treatment comprising the steps of:
- 20 (a) administering to said animal an augmenting agent which enhances CREB pathway function; and

- (b) training said animal under conditions sufficient to produce an improvement in performance by said animal of a cognitive task whose deficit is associated with said cerebrovascular disease, whereby said cognitive deficit is treated.
- 5 50. The method of Claim 49 wherein a performance gain is achieved relative to the performance of said cognitive task achieved by training alone.
  - 51. The method of Claim 49 wherein said cerebrovascular disease is selected from the group consisting of stroke and ischemia.
- 52. The method of Claim 49 wherein in step b), training comprises multiple training sessions.
  - 53. The method of Claim 49 wherein said augmenting agent is administered before and during each training session.
  - 54. The method of Claim 49 wherein said animal is a mammal.
  - 55. The method of Claim 54 wherein said mammal is a human.
- 15 56. The method of Claim 49 wherein said augmenting agent induces CREB-dependent gene expression.
  - 57. A method of treating a cognitive deficit associated with a trauma dependent loss of cognitive function in an animal in need of said treatment comprising the steps of:
- 20 (a) administering to said animal an augmenting agent which enhances CREB pathway function; and

- (b) training said animal under conditions sufficient to produce an improvement in performance by said animal of a cognitive task whose deficit is associated with said trauma dependent loss of cognitive function,
- 5 whereby said cognitive deficit is treated.
  - 58. The method of Claim 57 wherein a performance gain is achieved relative to the performance of said cognitive task achieved by training alone.
  - 59. The method of Claim 57 wherein said trauma dependent loss of function is selected from the group consisting of: head trauma and brain trauma.
- 10 60. The method of Claim 57 wherein in step b), training comprises multiple training sessions.
  - 61. The method of Claim 60 wherein said augmenting agent is administered before and during each training session.
  - 62. The method of Claim 57 wherein said animal is a mammal.
- 15 63. The method of Claim 62 wherein said mammal is a human.
  - 64. The method of Claim 57 wherein said augmenting agent induces CREB-dependent gene expression.
  - 65. A method of treating a cognitive deficit associated with a genetic defect in an animal in need of said treatment comprising the steps of:
- 20 (a) administering to said animal an augmenting agent which enhances CREB pathway function; and

- (b) training said animal under conditions sufficient to produce an improvement in performance by said animal of a cognitive task associated with said genetic defect,
- whereby said cognitive deficit is treated.
- 5 66. The method of Claim 65 wherein a performance gain is achieved relative to the performance of said cognitive task achieved by training alone.
  - 67. The method of Claim 65 wherein said genetic defect is selected from the group consisting of: Rubinstein-Taybi syndrome and down syndrome.
- 68. The method of Claim 65 wherein in step b), training comprises multiple training sessions.
  - 69. The method of Claim 68 wherein said augmenting agent is administered before and during each training session.
  - 70. The method of Claim 65 wherein said animal is a mammal.
  - 71. The method of Claim 70 wherein said mammal is a human.
- 15 72. The method of Claim 65 wherein said augmenting agent induces CREB-dependent gene expression.
  - 73. A method of improving learning in an animal with a learning disability comprising the steps of:
- (a) administering to said animal an augmenting agent which enhances CREB pathway function; and

- (b) training said animal under conditions sufficient to produce an improvement in performance by said animal of a cognitive task whose deficit is associated with said learning disability, whereby learning is improved.
- 5 74. The method of Claim 73 wherein a performance gain is achieved relative to the performance of said cognitive task achieved by training alone.
  - 75. The method of Claim 73 wherein in step b), training comprises multiple training sessions.
- 76. The method of Claim 75 wherein said augmenting agent is administered before and during each training session.
  - 77. The method of Claim 73 wherein said animal is a mammal.
  - 78. The method of Claim 77 wherein said mammal is a human.
  - 79. The method of Claim 73 wherein said augmenting agent induces CREB-dependent gene expression.
- 15 80. A method for repeated stimulation of neuronal activity or a pattern of neuronal activity in an animal comprising the steps of:
  - (a) administering to said animal an augmenting agent which enhances CREB pathway function; and
  - (b) training said animal under conditions sufficient to stimulate neuronal activity or a pattern of neuronal activity in said animal.

- 81. The method of Claim 80 wherein in step b), training comprises multiple training sessions.
- 82. The method of Claim 81 wherein said augmenting agent is administered before and during each training session.
- 5 83. The method of Claim 80 wherein said animal is a mammal.
  - 84. The method of Claim 83 wherein said mammal is a human.
  - 85. The method of Claim 80 wherein said augmenting agent induces CREB-dependent gene expression.
- 86. A method of therapy of a cognitive deficit associated with a central nervous

  system disorder or condition in an animal having undergone neuronal stem cell
  manipulation comprising the steps of:
  - (a) administering to said animal an augmenting agent which enhances CREB pathway function; and
  - (b) training said animal under conditions sufficient to stimulate neuronal activity or a pattern of neuronal activity in said animal.
  - 87. The method of Claim 86 wherein training in step b) further produces an improvement in performance by said animal of a cognitive task whose deficit is associated with said central nervous system disorder or condition.
- The method of Claim 87 wherein a performance gain is achieved relative to the performance of said cognitive task achieved by training alone.

- 89. The method of Claim 86 wherein in step b), training comprises multiple training sessions.
- 90. The method of Claim 89 wherein said augmenting agent is administered before and during each training session.
- 5 91. The method of Claim 86 wherein said animal is a mammal.
  - 92. The method of Claim 91 wherein said animal is a human.
  - 93. The method of Claim 86 wherein said augmenting agent induces CREB-dependent gene expression.